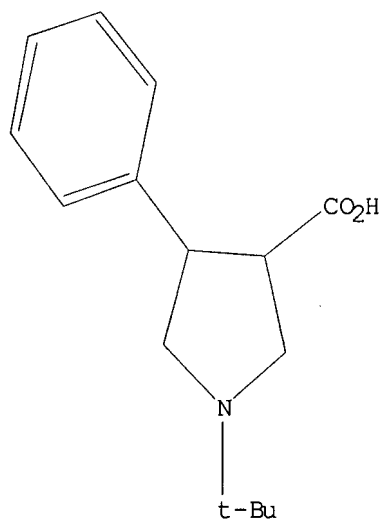


10/550,640

STN structure
Search

05/30/2007

(Registry / Caplus)



G1 H, MeO

Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 15:56:25 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 47 TO ITERATE

100.0% PROCESSED 47 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 529 TO 1351

PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s l1 full

FULL SEARCH INITIATED 15:56:30 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 994 TO ITERATE

100.0% PROCESSED 994 ITERATIONS

SEARCH TIME: 00.00.01

8 ANSWERS

L3 8 SEA SSS FUL L1

=> fil caplus

COST IN U.S. DOLLARS

SINCE FILE

ENTRY

TOTAL

SESSION

FULL ESTIMATED COST

172.10

172.31

FILE 'CAPLUS' ENTERED AT 15:56:36 ON 30 MAY 2007

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PLEASE SEE "HELP USAGETERMS" FOR DETAILS.

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Searched by Jason M. Nolan, Ph.D.

05/30/2007

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FILE COVERS 1907 - 30 May 2007 VOL 146 ISS 23
FILE LAST UPDATED: 29 May 2007 (20070529/ED)

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<http://www.cas.org/infopolicy.html>

=> s 13
L4

14 L3

=> d ibib abs hitstr 1-14

L4 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2007:410486 CAPLUS
 DOCUMENT NUMBER: 146:421856
 TITLE: Acylated spiropiperidine derivatives as
 melanocortin-4

receptor modulators and their preparation,
 pharmaceutical compositions and use in the treatment
 of diseases

INVENTOR(S): Jian, Tianying; Liu, Jian; Nargund, Ravi P.

PATENT ASSIGNEE(S): Merck & Co., Inc., USA

SOURCE: PCT Int. Appl., 159pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2007041052	A2	20070412	WO 2006-US37196	20060925
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW			
RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			

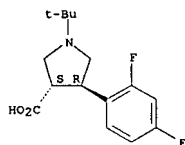
PRIORITY APPLN. INFO.: US 2005-721678P P 20050929

GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Certain N-acylated spiropiperidine derivs. of formula I are ligands of the human melanocortin receptor(s) and, in particular, are selective ligands of the human melanocortin-4 receptor (MC4R). They are therefore useful for the treatment, control, or prevention of diseases and disorders responsive to the modulation of MC4R, such as obesity, diabetes, nicotine addiction, alcoholism, sexual dysfunction, including erectile dysfunction and female sexual dysfunction. Comps. of formula I wherein A is (un)substituted heteroaryl; X and Y taken together to form CR6=CR6; one of X and Y is CR62 and the other is CR62, NR6, CO C=NR6, C=CR62, O, S, SO and SO2; one of X and Y is NR9, and the other is CR62, CO, C=NR6, C=CR62, SO and SO2; one of X and Y is CO and the other is CR62, NR6, C=NR6, O and S;

L4 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



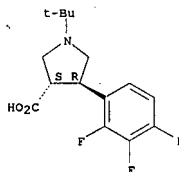
L4 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

Z is CH and N, provided that when Z is N, R1 is not NH2 and derivs.: R1 is H, (alkyl)amine, amidino, C1-6 alkyl, (hetero)aryl, (alkyl)(hetero)aryl, etc.; R2 is Ph, naphthyl and heteroaryl; R6 is H, C1-6 alkyl, (alkyl)(hetero)aryl, alkoxyalkyl, cyanoalkyl, etc.; R9 is H, C1-6 alkyl, (alkyl)(hetero)aryl, (alkyl)-C3-7 cycloalkyl, alkylcarboxylic acid derivs., etc.; R11 is H, OH, C1-8 alkyl, O-C1-8 alkyl, halo, NH and derivs., SH and derivs., and CF3; m and n are independently 1 and 2; and their pharmaceutically acceptable salts thereof, are claimed. Example compd.

II was prepd. by a multistep procedure (procedure given). All the invention compds. were evaluated for their melanocortin-4 receptor modulatory activity. The tested compds. were found to bind to melanocortin-4 receptor with IC50 values less than 10 µM.

IT 934361-22-5P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (intermediate; preparation of acylated spiropiperidine derivs. as melanocortin-4 receptor modulators)
 RN 934361-22-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 1-(1,1-dimethylethyl)-4-(2,3,4-trifluorophenyl)-, (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry.



IT 455957-94-5
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (starting material; preparation of acylated spiropiperidine derivs. as melanocortin-4 receptor modulators)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

L4 ANSWER 1 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

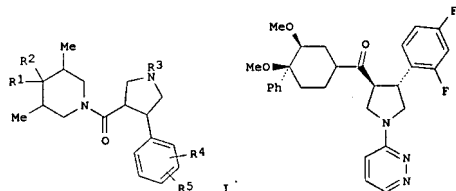
L4 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2007:150961 CAPLUS
 DOCUMENT NUMBER: 146:229190
 TITLE: Preparation of pyrrolidinylcarbonylpiperidines as melanocortin subtype 4 (MC4) receptor agonists for treatment of lower urinary tract dysfunction.
 INVENTOR(S): McMurray, Gordon; Phillips, Stephen Charles; Westbrook, Simon Lempriere
 PATENT ASSIGNEE(S): Pfizer Limited, UK
 SOURCE: PCT Int. Appl., 85pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2007015157	A2	20070208	WO 2006-1B2119	20060720
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW			
RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			

PRIORITY APPLN. INFO.: GB 2005-15817 A 20050801
 US 2005-705237P P 20050802

OTHER SOURCE(S): MARPAT 146:229190
 GI



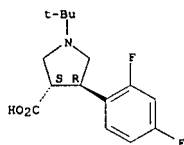
AB Use of MC4 receptor antagonists, e.g. {I; R1 = (substituted) alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, aryl, heterocyclyl, etc.: R2

L4 ANSWER 2 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
H, OH, OMe; R3 = H, R4, R5 = H, halo, alkyl, alkenyl, alkynyl, cyano,
CF3, CH2CF3, OCF3, OCH2CF3, cycloalkenyl(alkyl), amino(alkyl), etc.; R4R5
= atoms to form fused 5-7 membered (unsatd.) ring] for the manuf. of a
medicament for the treatment of lower urinary tract dysfunction is
claimed. Thus, title compd. (II) (prepn. outlined) at 2 mg/kg i.v. in

the
dog urethral pressure model gave a 20% increase in peak urethral
pressure.
IT 455957-94-5P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)
(preparation of pyrrolidinylcarbonylpiperidines as MC4 receptor
agonists for
treatment of lower urinary tract dysfunction)

RN 455957-94-5 CAPLUS
CN 3-Pyrrolidinecarboxylic acid,
4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-
, (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

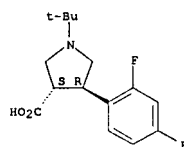


L4 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
HCl afforded compd. II·HCl. Comps. according to the invention
herein exhibited a binding const. at the MCR4 (melanocortin receptor 4)
expressed as an Ki value against AGRP (agouti related protein) lower than
1000 nM. Comps. I are claimed useful for the treatment of sexual
dysfunction and obesity.

IT 455957-75-2P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
(Reactant or reagent)
(preparation of piperidinoyl-pyrrolidine and piperidinoyl-piperidine
comps.
as MCR4 agonists)

RN 455957-75-2 CAPLUS
CN 3-Pyrrolidinecarboxylic acid,
4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-
, hydrochloride (1:1), (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



● HCl

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS
RECORD. ALL CITATIONS AVAILABLE IN THE RE

FORMAT

L4 ANSWER 3 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 2007:150182 CAPLUS
DOCUMENT NUMBER: 146:229186
TITLE: Preparation of piperidinoyl-pyrrolidine and
piperidinoyl-piperidine compounds as MCR4 agonists
Inventors, Mark David; Brown, Alan Daniel; Fradet,
David
PATENT ASSIGNEE(S): Sebastian; Lansdell, Mark Ian
Pfizer Limited, UK
SOURCE: PCT Int. Appl., 103pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2007015162	A1	20070208	WO 2006-1B2151	20060726
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MM, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW			
RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
PRIORITY APPLN. INFO.:			US 2005-706191P	P 20050804

OTHER SOURCE(S): MARPAT 146:229186
GI

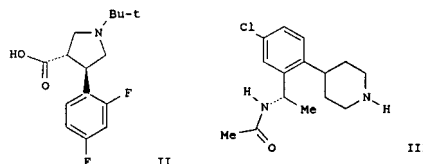
* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Title compds. I [n = 1, 2; R6 = H, alkyl, cycloalkyl, etc.; R7 =
pyridinyl
or Ph (wherein pyridinyl or Ph is optionally substituted by halo, CN,
CF3,
etc.); R10 = Q1; R1, R4 = H, alkyl, OH, etc.; R2 = H, OH, O-alkyl, etc.;
R3 = aryl or heteroaryl (wherein said moieties are optionally substituted
by halo, CN, CF3, etc.); R5 = H or alkyl; R1, R4 and R5 are not all
simultaneously H.; when R1 is Me and R4 is H, then R5 is not Me.; when R4
is Me and R5 is H, then R1 is not Me.; when R5 is Me and R4 is H, then R1
is not Me.], pharmaceutically acceptable salts, hydrates, solvates,
polymorphs, and prodrugs thereof were prepared. For example, amidation of
(3S,4R)-1-tert-butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid
hydrochloride, e.g., prepared from (E)-2,4-difluorocinnamic acid in 3
steps,
with (3S,4R)-3-methyl-4-phenylpiperidin-4-ol followed by treatment with

L4 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER: 2007:14419 CAPLUS
DOCUMENT NUMBER: 146:121832
TITLE: Preparation of pyrrolidinylcarbonylpiperidine
derivative as MC4R agonist and of crystalline forms
of
said derivative
Inventors, Calabria, Ralph; Cheng, Yu; Ferlita, Russell R.;
Kamali, Ashkan; Murry, Jerry A.; Mathre, David;
Peressypkin, Andrey V.; Thompson, Karen; Wang, Jian;
Wenslow, Robert M.
PATENT ASSIGNEE(S): Merck & Co., Inc., USA
SOURCE: PCT Int. Appl., 50pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2007002462	A2	20070104	WO 2006-US24573	20060623
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MM, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SI, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW			
RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
PRIORITY APPLN. INFO.:			US 2005-694488P	P 20050627

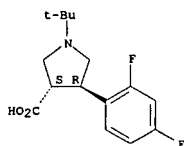
GI



AB The present invention relates to a process for producing

N-((1S)-1-[2-(1-((3S,4R)-1-tert-butyl-4-(2,4-difluorophenyl)pyrrolidin-3-yl)carbonyl)piperidin-4-yl]-5-chlorophenyl)ethyl)acetamide (II), and novel

L4 ANSWER 4 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
 cryst. salts, hydrates, solvates, and polymorphic forms thereof. Thus, treatment of pyrrolidinecarboxylic acid II with CDI in acetonitrile, followed by reaction with piperidine deriv. III in acetonitrile in the presence of triethylamine gave, after workup, I (free base). A soln. of I (free base) in iso-Pr acetate was mixed with 0.5 N HCl/H₂O soln. to give, after workup and crystn., the cryst. hydrate mono-HCl salt of I.
 IT 455957-94-5
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (preparation of pyrrolidinylcarbonylpiperidine derivative as MC4R agonist and of crystalline forms of said derivative)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).



L4 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
 ACCESSION NUMBER: 2006:165649 CAPLUS
 DOCUMENT NUMBER: 144:232924
 TITLE: Preparation of piperidine derivatives as melanocortin-4 receptor agonists
 INVENTOR(S): Barakat, Khaled J.; Guo, Liangqin; Liu, Jian; Nargund, Ravi P.; Sehbat, Iyassu K.; Ye, Zhixiong
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: FCT Int. Appl., 75 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

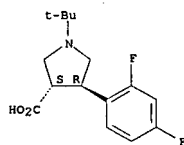
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006019787	A2	20060223	WO 2005-US24806	20050713
WO 2006019787	A3	20060914		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	AF, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
AU 2005275232	A1	20060223	AU 2005-275232	20050713
CA 2573714	A1	20060223	CA 2005-2573714	20050713
EP 1773347	A2	20070418	EP 2005-769526	20050713
PRIORITY APPLN. INFO.:			US 2004-588494P	P 20040716
			WO 2005-US24806	W 20050713

OTHER SOURCE(S): MARPAT 144:232924
 GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The title piperidine derivs. I [wherein m = 0-2; n = 1-2; R1 and R2 = independently halo, CF₃, CH₃, and OMe; R3 and R4 = independently halo, CF₃, CN, alkyl, alkoxy, etc.; R5 = OH, halo, alkyl, alkoxy, etc.], or pharmaceutically acceptable salts thereof were prepared as agonists of the human melanocortin-4 receptors (MCR-4). For example, II was prepared in a multi-step synthesis. The title compds. showed IC₅₀ less than 10 μ M against MCR-4. Formulations with finely divided lactose as hard gelatin

L4 ANSWER 5 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
 capsule have been described. The compds. are useful for the treatment, control, or prevention of diseases and disorders responsive to the activation of MCR-4, such as obesity, diabetes, male or female sexual dysfunction.
 IT 455957-94-5P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (intermediate; preparation of piperidine derivs. as MCR-4 agonists)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).

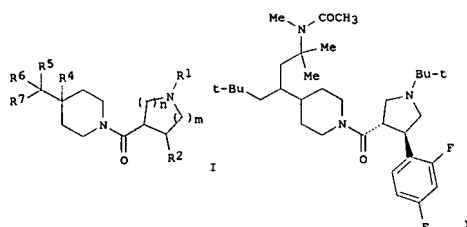


L4 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
 ACCESSION NUMBER: 2006:164650 CAPLUS
 DOCUMENT NUMBER: 144:254006
 TITLE: Preparation of piperidine derivatives as melanocortin-4 receptor agonists
 INVENTOR(S): Bakshi, Raman K.; Dellureficio, James P.; Nargund, Ravi P.
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: FCT Int. Appl., 78 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006020277	A2	20060223	WO 2005-US25505	20050715
WO 2006020277	A3	20060720		
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	AF, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
AU 2005274701	A1	20060223	AU 2005-274701	20050715
CA 2574156	A1	20060223	CA 2005-2574156	20050715
EP 1773338	A2	20070418	EP 2005-773009	20050715
R:	AF, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR			
PRIORITY APPLN. INFO.:			US 2004-589089P	P 20040719
			WO 2005-US25505	W 20050715

OTHER SOURCE(S): MARPAT 144:254006
 GI

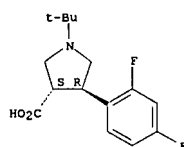
L4 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN (Continued)



AB The title piperidine derivs. I [wherein m = 0-2; n = 1-2; R1 = H, amino, alkyl, etc.; R2 = (un)substituted Ph, naphthyl, or heteroaryl; R4 = H, alkyl, alkoxy, etc.; R5 = CF3, alkyl, alkenyl, alkynyl, etc.; R6 = H, alkyl, or alkoxy; R7 = NH2, CN, OH, alkoxy, etc.], or pharmaceutically acceptable salts thereof were prepared as agonists of the human melanocortin-4 receptors (MCR-4). For example, II was prepared in a multi-step synthesis. The title compds. showed IC50 less than 10 μ M against MCR-4. Formulations with finely divided lactose as hard gelatin capsule have been described. The compds. are useful for the treatment, control, or prevention of diseases and disorders responsive to the activation of MCR-4, such as obesity, diabetes, male or female sexual dysfunction (no data).

IT 455957-94-5P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(intermediate; preparation of piperidine derivs. as MCR-4 agonists)
RN 455957-94-5 CAPLUS
CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L4 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN

ACCESSION NUMBER: 2005:715328 CAPLUS
DOCUMENT NUMBER: 143:211844
TITLE: Preparation of Pyrrolidinyl(carbonyl) piperidines as melanocortin receptor 4 agonists for therapeutic use
INVENTOR(S): Calabrese, Andrew Antony; Fradet, David Sebastian; Hepworth, David; Lansdell, Mark
PATENT ASSIGNEE(S): Pfizer Inc, UK
SOURCE: U.S. Pat. Appl. Publ., 102 pp.
CODEN: USXXCO
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

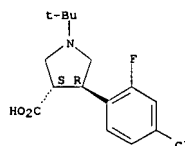
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2005176772	A1	20050811	US 2005-51144	20050203
AU 2005213538	A1	20050825	AU 2005-213538	20050126
CA 2555800	A1	20050825	CA 2005-2555800	20050126
WO 2005077935	A1	20050825	WO 2005-1B208	20050126
W: AE, AG, AL, AM, AT, AU, AZ, BA, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MY, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, ME, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, BG, BR, BU, BU, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
EP 1716135	A1	20061102	EP 2005-702363	20050126
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, BA, HR, IS, YU				
CN 1930193	A	20070314	CN 2005-80007573	20050126
NL 1028193	A1	20050808	NL 2005-1028193	20050204
NL 1028193	C2	20060420		
NO 2006003944	A	20061031	NO 2006-3944	20060904
PRIORITY APPLN. INFO.:			GB 2004-2492	A 20040204
			US 2004-543950P	P 20040212
			GB 2004-12417	A 20040603
			US 2004-580209P	P 20040615
			GB 2004-25530	A 20041119
			US 2004-634916P	P 20041210
			WO 2005-1B208	W 20050126

OTHER SOURCE(S): MARPAT 143:211844
GI

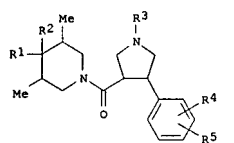
L4 ANSWER 6 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN (Continued)

IT 757974-04-2
RL: RCT (Reactant); RACT (Reactant or reagent)
(preparation of piperidine derivs. as MCR-4 agonists)
RN 757974-04-2 CAPLUS
CN 3-Pyrrolidinecarboxylic acid, 4-(4-chloro-2-fluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L4 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN (Continued)

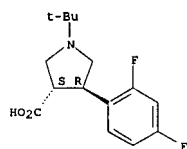


AB The present invention relates to a class of melanocortin MCR4 agonists of general formula (I) wherein R1, R2, R3, R4 and R5 are as defined below and especially to selective MCR4 agonist compds., to their use in medicine, to compns. containing them, to processes for their preparation and to intermediates used in such processes. The variables for I are: R1 = optionally substituted alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, alkylcycloalkyl, aryl, alkylaryl, heterocycle, or alkylheterocycle; R2 = H, OH or OCH3; R3 = H, or optionally substituted alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, alkylcycloalkyl, aryl, alkylaryl, heterocycle, or alkylheterocycle; R4 = H, alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, halogen, (CH2)pOR6, (CH2)pNR7R8, CN, C(O)R6, C(O)OR6, C(O)NR7R8, (CH2)pNR7SO2R8, CF3, CH2CF3, OCF3 or OCH2CF3 wherein p=0, 1 or 2; R5 = alkyl, alkenyl, alkynyl, cycloalkyl, cycloalkenyl, halogen, (CH2)pOR6, (CH2)pNR7R8, CN, C(O)R6, C(O)OR6, C(O)NR7R8, (CH2)pNR7SO2R8, CF3, CH2CF3, OCF3 or OCH2CF3 wherein p=0, 1 or 2; or R4 and R5 form a fused ring; R6, R7 and R8 = H, CH3 or CH2CH3. Compds. of the present invention are particularly suitable for treating female sexual dysfunction, male erectile dysfunction, obesity and diabetes.

IT 455957-75-2P, (3S,4R)-1-tert-Butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid hydrochloride
455957-94-5P, (3S,4R)-1-tert-Butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid 862282-17-5P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of Pyrrolidinylcarbonyl piperidines as melanocortin receptor 4 agonists for therapeutic use)
RN 455957-75-2 CAPLUS
CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, hydrochloride (1:1), (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

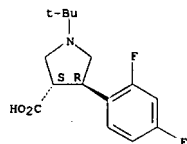
L4 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



● HCl

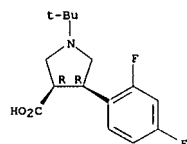
RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid,
 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-
 , (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



RN 862282-17-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid,
 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-
 , (3R,4R)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



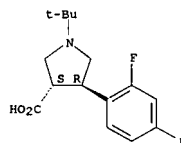
L4 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:283875 CAPLUS
 DOCUMENT NUMBER: 143:7543
 TITLE: Enantioselective Nitrile Anion Cyclization to Substituted Pyrrolidines. A Highly Efficient
 Synthesis of
 (3S,4R)-N-tert-Butyl-4-Arylpyrrolidine-3-Carboxylic Acid
 AUTHOR(S): Chung, John Y. L.; Cvetovich, Raymond; Amato, Joseph; McWilliams, J. Christopher; Reamer, Robert;
 DiMichele, Lisa
 CORPORATE SOURCE: Department of Process Research, Merck Research Laboratories, Rahway, NJ, 07065, USA
 SOURCE: Journal of Organic Chemistry (2005), 70(9), 3592-3601
 CODEN: JOCEAH; ISSN: 0022-3263
 PUBLISHER: American Chemical Society
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 OTHER SOURCE(S): CASREACT 143:7543
 AB A practical asym. synthesis of N-tert-Bu disubstituted pyrrolidines via a nitrile anion cyclization strategy is described. The five-step chromatog.-free synthesis of (3S,4R)-1-tert-butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid from 2-chloro-1-(2,4-difluorophenyl)ethanone achieved a 71% overall yield. The cyclization substrate was prepared via a catalytic CBS asym. reduction, tert.-butylamine displacement of the chlorohydrin, and a conjugate addition of the hindered secondary amine to acrylonitrile. The key nitrile anion 5-exo-tet cyclization concomitantly formed the pyrrolidine ring with clean inversion of the C-4 center to afford 1,3,4-trisubstituted chiral pyrrolidine in >95% yield and 94-99% ee. Di-Et chlorophosphate and lithium hexamethylsilazide were shown to be the resp. optimum activating group and base in this cyclization. The trans-cis mixture of the pyrrolidine nitrile undergoes a kinetically controlled epimerization/ saponification to afford the pure trans-pyrrolidine carboxylic acid target compound in >99.9% chemical and optical purity. This chemical was also shown to be applicable to both electronically neutral and rich substituted Ph substrates.
 IT 455957-94-5P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (preparation of (3S,4R)-N-tert-butyl-4-arylpyrrolidine-3-carboxylic acids by enantioselective nitrile anion cyclization)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid,
 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-
 , (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

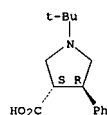
L4 ANSWER 7 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

L4 ANSWER 8 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



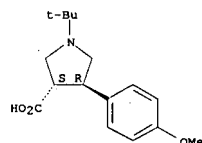
IT 782482-56-8P 782482-57-9P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (preparation of (3S,4R)-N-tert-butyl-4-arylpyrrolidine-3-carboxylic acids by enantioselective nitrile anion cyclization)
 RN 782482-56-8 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 1-(1,1-dimethylethyl)-4-phenyl-, (3S,4R)-
 (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 782482-57-9 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 1-(1,1-dimethylethyl)-4-(4-methoxyphenyl)-,
 (3S,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



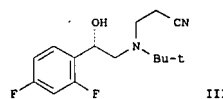
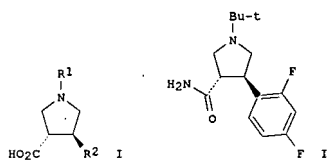
REFERENCE COUNT: 51 THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

L4 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
 ACCESSION NUMBER: 2004:902344 CAPLUS
 DOCUMENT NUMBER: 141:379798
 TITLE: A process for preparation of pyrrolidinecarboxylic acid derivatives, useful for treatment of melanocortin receptor mediated diseases
 INVENTOR(S): Ovetovich, Raymond; Chung, John Y.; Amato, Joseph S.; Dimichele, Lisa
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: PCT Int. Appl., 45 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
 WO 2004092126 A2 20041028 WO 2004-US11253 20040409
 WO 2004092126 A3 20050120
 WO 2004092126 B1 20050331
 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
 CA 2521487 A1 20041028 CA 2004-2521487 20040409
 EP 1615882 A2 20060118 EP 2004-750027 20040409
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK
 HR CN 1774419 A 20060517 CN 2004-80009882 20040409
 JP 2006523700 T 20061019 JP 2006-509940 20040409
 US 2006199958 A1 20060907 US 2005-550640 20050923
 US 2003-462795P P 20030414
 PRIORITY APPLN. INFO.:
 WO 2004-US11253 W 20040409
 OTHER SOURCE(S): CASREACT 141:379798; MARPAT 141:379798
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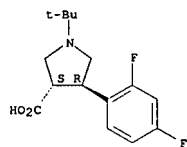
Instant App

L4 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

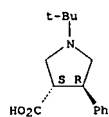


AB The invention relates to a preparation of pyrrolidinecarboxylic acid derivs. of formula I [wherein: R1 is H, NH2, alkyl, or (CH2)0-4-heteroaryl, etc.; R2 is (cyclo)alkyl, furyl, oxazolyl, or thienyl, etc.]. These compds. are useful for treatment of melanocortin receptor mediated diseases such as obesity, diabetes, male sexual dysfunction, and female sexual dysfunction (no biol. data). For instance, pyrrolidinecarboxylic acid derivative II was prepared via intramol. heterocyclization of the obtained amine III and hydrolysis (example 1).
 IT RL: IMF (Industrial manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (process for preparation of pyrrolidinecarboxylic acid derivs. useful for treatment of melanocortin receptor mediated diseases)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 1-(1,1-dimethylethyl)-4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).

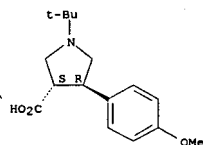
L4 ANSWER 9 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



IT 782482-56-8P 782482-57-9P
 RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
 for (process for preparation of pyrrolidinecarboxylic acid derivs. useful for treatment of melanocortin receptor mediated diseases)
 RN 782482-56-8 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 1-(1,1-dimethylethyl)-4-phenyl-, (3S,4R)- (9CI) (CA INDEX NAME)
 Absolute stereochemistry.



RN 782482-57-9 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 1-(1,1-dimethylethyl)-4-(4-methoxyphenyl)-, (3S,4R)- (9CI) (CA INDEX NAME)
 Absolute stereochemistry.



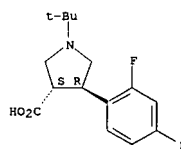
L4 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
 ACCESSION NUMBER: 2004:872662 CAPLUS
 DOCUMENT NUMBER: 141:366128
 TITLE: Preparation of cycloalkylcarbonyl or heterocycloalkylcarbonyl-substituted spiropiperidines as melanocortin-4 receptor agonists for the treatment of conditions such as obesity
 INVENTOR(S): Guo, Liangqin; He, Shuwen; Jian, Tianying; Lai, Yingjie; Liu, Jian; Nargund, Ravi P.; Sebbat, Iyassu K.; Ujjainwalla, Perce; Ye, Zhixiong; Young, Jonathan
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: PCT Int. Appl., 200 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:
 PATENT NO. KIND DATE APPLICATION NO. DATE
 WO 2004089307 A2 20041021 WO 2004-US9751 20040331
 WO 2004089307 A3 20050331
 WO 2004089307 A8 20050623
 W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
 RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
 AU 2004227835 A1 20041021 AU 2004-227835 20040331
 CA 2520114 A1 20041021 CA 2004-2520114 20040331
 EP 1613601 A2 20060111 EP 2004-749540 20040331
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 BR 2004009078 A 20060418 BR 2004-9078 20040331
 CN 1768041 A 20060503 CN 2004-80009148 20040331
 JP 2006522132 T 20060928 JP 2006-509489 20040331
 JP 3856815 B2 20061213
 US 2006183904 A1 20060817 US 2005-548350 20050907
 NO 2005005166 A 20051230 NO 2005-5166 20051103
 PRIORITY APPLN. INFO.: US 2003-460293P P 20030404
 WO 2004-US9751 W 20040331
 OTHER SOURCE(S): MARPAT 141:366128
 GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Title compds. I or II [X,Y = R62C, R9N, C(=O); Y,X = R62C, R6N, C(=O)].

L4 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
 R6N:C, O, S, S(:O), SO2; XY = CR6:CR6; Z = R1C, N; A = (CH2)m; E = (CH2)p;
 R1 = H, amidino, (un)substituted aminoalkyl, iminoalkyl, alkyl, cycloalkylalkyl, phenylalkyl, naphthylalkyl, or heteroarylalkyl; R2 = (un)substituted Ph, naphthyl, heteroaryl; R4 = H, (un)substituted alkyl, halogen, alkoxy, O2N, F3C, F3CCH2, F3CO, F3CCH2O; R6, R9 = H, (un)substituted alkyl, phenylalkyl, naphthylalkyl, heteroarylalkyl, cycloalkylalkyl, heterocycloalkylalkyl, aminoalkyl, carboxyalkyl, etc.; m, p = 1, 2; n = 0-3] such as III-HCl are prepd. as melanocortin-4 receptor agonists for the treatment of obesity and related conditions such as diabetes, bulimia, insulin resistance, and hyperlipidemia; a variety of other conditions, particularly male and female sexual dysfunction and erectile dysfunction, are also potentially treatable with the title compds. Oxoindanospiriperidinecarboxylate IV is reduced with sodium borohydride and the alc. eliminated in the presence of p-toluenesulfonic acid to give the indenespiroperidinecarboxylate; Jacobsen epoxidn. of the indene double bond, opening of the epoxide with sodium azide, aziridine formation using a fluorous phosphine, N-methylation of the aziridine, regioselective redn. of the aziridine with sodium borohydride to yield the aminoindanospiriperidinecarboxylate, acylation with 2-acetoxyisobutyl chloride, hydrolysis of the ester with sodium methoxide and methylation of the alc. with Me iodide, deprotection of the piperidine nitrogen, and acylation with nonracemic trans-4-(2,4-difluorophenyl)-1-tert-butyl-3-pyrrolidinecarboxylic acid yields III. Some of the title compds. bind to the melanocortin-4 receptor with IC50 values of <10 μ M and <5 μ M (no data).
 IT 455957-75-2P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (intermediate; preparation of cycloalkylcarbonyl or heterocycloalkylcarbonyl-substituted spiroperidines as melanocortin-4 receptor agonists for the treatment of conditions such as obesity and male or female sexual dysfunction)
 RN 455957-75-2 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, hydrochloride (1:1), (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).

L4 ANSWER 10 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

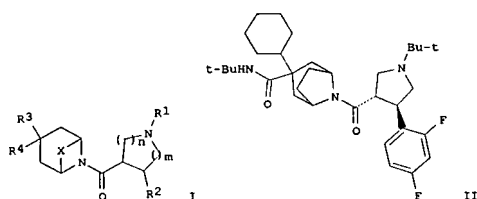


• HCl

L4 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
 ACCESSION NUMBER: 2004:857404 CAPLUS
 DOCUMENT NUMBER: 141:350043
 TITLE: Preparation of bicyclic piperidine derivatives as melanocortin-4 receptor agonists
 INVENTOR(S): Nargund, Ravi P.; Ye, Zhixiong; Guo, Liangqin
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: PCT Int. Appl., 103 pp.
 CODEN: PIXX2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004087159	A1	20041014	WO 2004-US8711	20040322
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GR, GU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
AU 2004226452	A1	20041014	AU 2004-226452	20040322
CA 2519939	A1	20041014	CA 2004-2519939	20040322
EP 1610789	A1	20060104	EP 2004-758182	20040322
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK				
CN 1764458	A	20060426	CN 2004-80008218	20040322
JP 2006521359	T	20060921	JP 2006-507449	20040322
US 2006194780	A1	20060931	US 2005-548762	20050912
PRIORITY APPLN. INFO.:			US 2003-457679P	P 20030326
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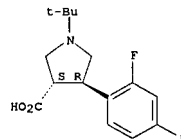
 OTHER SOURCE(S): MARPAT 141:350043
 GI



II

L4 ANSWER 11 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)

AB Title compds. I (R1 = H, amidino, alkyl, (un)substituted-arylalkyl, etc.; R2 = alkyl, (un)substituted-Ph, -naphthyl, and -heteroaryl; R3 and R4 independently = H, alkyl, alkenyl, cyanoalkyl, etc.; X = (CR5R6)p where R5 and R6 independently = H, alkyl, OH, amino, halo, cycloalkylalkyl, etc. and p = 1-3; m = 0-2; n = 1-2) are prepared and disclosed as agonists of the human melanocortin receptor(s) and, in particular, are selective agonists of the human melanocortin-4 receptor (MC-4R). Thus, e.g., II was prepared in a multistep synthesis from N-carbethoxy-4-tropinone to provide 3-[(tert-butylamino)carbonyl]-3-cyclohexyl-8-azabicyclo[3.2.1]octane which was amidated with (3R,4S)-1-tert-butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid (preparation given). Representative compds. of the invention were tested and found to bind to the melanocortin-4 receptor and possessed IC50 values less than 10 μ M. They are therefore useful for the treatment, control, or prevention of diseases and disorders responsive to the activation of MC-4R, such as obesity, diabetes, sexual dysfunction, including erectile dysfunction and female sexual dysfunction.
 IT 455957-94-5
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (starting material; preparation of bicyclic piperidine deriva. as melanocortin-4 receptor agonists)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE
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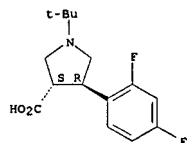
L4 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN
 ACCESSION NUMBER: 2004:756685 CAPLUS
 DOCUMENT NUMBER: 141:277640
 TITLE: Preparation of acylated piperazine derivatives as melanocortin-4 receptor agonists for the treatment of obesity, diabetes mellitus and sexual dysfunction, and pharmaceutical compositions thereof
 INVENTOR(S): Bakshi, Raman K.; Guo, Liangqin; Hong, Qingmei; Nargund, Ravi P.; Pollard, Patrick G.; Sehbat, Iyassu K.; Ujjainwalla, Feroze; Ye, Zhixiong
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: PCT Int. Appl., 187 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004078717	A1	20040916	WO 2004-US7713	20040227
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GR, GU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NA, NI, NG, NO, NZ, OM, OS, PA, PE, PG, PH, PK, PL, PT, QA, RO, RU, RW, SA, SC, SD, SE, SG, SI, SK, SL, SM, SN, SR, SS, ST, SV, SW, SY, SZ, TD, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VE, VU, WO, XA, XB, XC, XE, XF, XG, XH, XI, XJ, XK, XL, XM, XN, XO, XP, XQ, XR, XS, XT, XU, XV, XW, XX, XY, XZ, YD, YE, YF, YG, YH, YI, YJ, YK, YL, YM, YN, YO, YP, YQ, YR, YS, YT, YU, YV, YW, YY, YZ, ZA, ZB, ZC, ZD, ZE, ZF, ZG, ZH, ZI, ZJ, ZK, ZL, ZM, ZN, ZO, ZP, ZQ, ZR, ZS, ZT, ZU, ZV, ZW, ZX, ZY, ZZ			
RW:	BW, GH, GM, GR, GU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NA, NI, NG, NO, NZ, OM, OS, PA, PE, PG, PH, PK, PL, PT, QA, RO, RU, RW, SA, SC, SD, SE, SG, SI, SK, SL, SM, SN, SR, SS, ST, SV, SW, SY, SZ, TD, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VE, VU, WO, XA, XB, XC, XE, XF, XG, XH, XI, XJ, XK, XL, XM, XN, XO, XP, XQ, XR, XS, XT, XU, XV, XW, XX, XY, XZ, YD, YE, YF, YG, YH, YI, YJ, YK, YL, YM, YN, YO, YP, YQ, YR, YS, YT, YU, YV, YW, YY, YZ, ZA, ZB, ZC, ZD, ZE, ZF, ZG, ZH, ZI, ZJ, ZK, ZL, ZM, ZN, ZO, ZP, ZQ, ZR, ZS, ZT, ZU, ZV, ZW, ZX, ZY, ZZ			
US 2004204398	A1	20041014	US 2004-788859	20040227
US 7160886	B2	20070109		
PRIORITY APPLN. INFO.:			US 2003-451502P	P 20030303
			US 2003-515943P	P 20031030

OTHER SOURCE(S): MARPAT 141:277640
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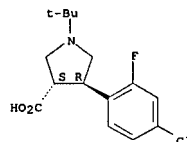
L4 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN (Continued)
 (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



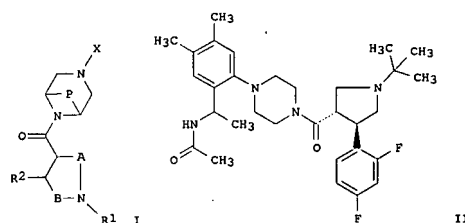
IT 757974-04-2
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (reactant; preparation of phenylpyrrolidinylcarbonyl/phenylpiperidinylcarbonyl piperazines as melanocortin-4 receptor agonists)
 RN 757974-04-2 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(4-chloro-2-fluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 3
 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RECORD
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L4 ANSWER 12 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN (Continued)

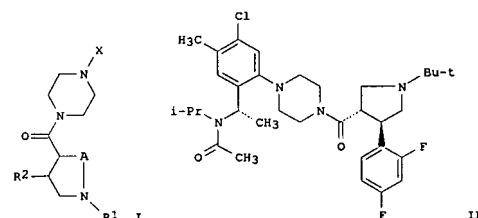


AB N-Acylated piperazine derivs. of formula I, wherein X is (un)substituted alkyl/phenyl/heteroaryl, etc.; R1 is H, amidino, (un)substituted (cyclo)alkyl/alkylimino/phenyl/naphthyl/heteroaryl, etc.; R2 is (un)substituted phenyl/naphthyl/heteroaryl; P is (CH2R4)m; R4 is H, (cyclo)alkyl/alkyl, aryl(alkyl), OH, halo, amino; m is 0-4; A is (CH2)r; r is 1 or 2; B = (CH2)s; s is 0-2, were prepared as melanocortin-4 receptor (MC-4R) agonists. II is one of the given examples, which was synthesized in several steps via coupling of the corresponding monosubstituted piperazine (preparation given) with the corresponding pyrrolidinecarboxylic acid (preparation given). Comps. I, including some specific stereoisomers, pharmaceutically acceptable salts thereof, and pharmaceutical compns. (examples given) comprising I and pharmaceutically acceptable carriers, are claimed. Certain compns. among I are agonists of the human melanocortin receptor(s) and, in particular, are selective agonists of the human MC-4R. Representative compns. (no indication) were tested and found to bind to the melanocortin-4 receptor, generally having IC50 values less than 10 μM. Representative compns. (no indication) were also tested in the functional model assay, and found generally to activate the MC-4R with EC50 values less than 10 μM. Comps. I are therefore useful for the treatment, control, or prevention of diseases and disorders responsive to the activation of MC-4R, such as obesity, diabetes, sexual dysfunction, including erectile dysfunction and female sexual dysfunction.
 IT 455957-94-5P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (intermediate; preparation of phenylpyrrolidinylcarbonyl/phenylpiperidinylcarbonyl piperazines as melanocortin-4 receptor agonists)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-

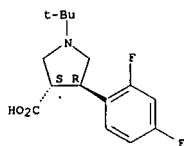
L4 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2007 ACS ON STN
 ACCESSION NUMBER: 2004:756684 CAPLUS
 DOCUMENT NUMBER: 141:277639
 TITLE: Preparation of acylated piperazine derivatives as melanocortin-4 receptor agonists for the treatment of obesity, diabetes mellitus and sexual dysfunction, and pharmaceutical compositions thereof
 INVENTOR(S): Bakshi, Raman K.; Hong, Qingmei; Nargund, Ravi P.; Pollard, Patrick G.; Sehbat, Iyassu K.; Ujjainwalla, Feroze; Ye, Zhixiong
 PATENT ASSIGNEE(S): Merck & Co. Inc., USA
 SOURCE: PCT Int. Appl., 76 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004078716	A1	20040916	WO 2004-US5982	20040227
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GR, GU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NA, NI, NG, NO, NZ, OM, OS, PA, PE, PG, PH, PK, PL, PT, QA, RO, RU, RW, SA, SC, SD, SE, SG, SI, SK, SL, SM, SN, SR, SS, ST, SV, SW, SY, SZ, TD, TH, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VE, VU, WO, XA, XB, XC, XE, XF, XG, XH, XI, XJ, XK, XL, XM, XN, XO, XP, XQ, XR, XS, XT, XU, XV, XW, XX, XY, XZ, YD, YE, YF, YG, YH, YI, YJ, YK, YL, YM, YN, YO, YP, YQ, YR, YS, YT, YU, YV, YW, YY, YZ, ZA, ZB, ZC, ZD, ZE, ZF, ZG, ZH, ZI, ZJ, ZK, ZL, ZM, ZN, ZO, ZP, ZQ, ZR, ZS, ZT, ZU, ZV, ZW, ZX, ZY, ZZ			
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US 2004204398	A1	20041014	US 2004-788859	20040227
US 7160886	B2	20070109		
PRIORITY APPLN. INFO.:			US 2003-451502P	P 20030303
			US 2003-515943P	P 20031030

OTHER SOURCE(S): MARPAT 141:277639
 GI



L4 ANSWER 13 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
 AB N-Acylated piperazine derivs. of formula I, wherein X is (un)substituted alkyl/phenyl/phenylalkyl; R1 is alkyl; R2 is (un)substituted phenyl; A is (CH2)_n; n is 1 or 2, or prepared as melanocortin-4 receptor (MC-4R) agonists. II is one of the given examples, which was synthesized in several steps via coupling of the left part, a phenylpiperazine (preparation given), with the right part, a phenylpyrrolidinecarboxylic acid (preparation given). Comps. I and pharmaceutically acceptable salts thereof as well as pharmaceutical compns. (examples given) comprising I and pharmaceutically acceptable carriers are claimed. Representative compds. (no indication) were tested and found to bind to the melanocortin-4 receptor, generally having IC50 values less than 10 μ M. Representative compds. (no indication) were also examined in several functional model tests, and found generally able to activate the MC-4R with EC50 values less than 10 μ M. Certain compds. are agonists of the human melanocortin receptor(s) and, in particular, are selective agonists of the human MC-4R. They are therefore useful for the treatment, control, or prevention of diseases and disorders responsive to the activation of MC-4R, such as obesity, diabetes, sexual dysfunction, including erectile dysfunction and female sexual dysfunction.
 IT 455957-94-SP
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (Intermediate; preparation of acylated piperazine derivs. as melanocortin-4 receptor agonists)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE
 FORMAT

L4 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN
 ACCESSION NUMBER: 2002:675993 CAPLUS
 DOCUMENT NUMBER: 137:216874
 TITLE: Acylated piperidine derivatives, specifically 1-(pyrrolidinylcarbonyl)piperidines, 1-(piperidinylcarbonyl)piperidines, and analogs, as melanocortin-4 receptor agonists, and their pharmaceutical compositions and therapeutic uses
 INVENTOR(S): Ujjainwalla, Feroze; Chu, Lin; Goulet, Mark T.; Lee, Bonnie; Warner, Daniel; Wyvrat, Matthew J.
 PATENT ASSIGNEE(S): Merck & Co., Inc., USA
 SOURCE: PCT Int. Appl., 112 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 2
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2002068388	A2	20020906	WO 2002-US5724	20020225
WO 2002068388	A3	20030313		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MX, MY, NZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW				
RW: GH, GM, KE, LS, MW, MZ, SD, SI, SZ, TZ, UG, ZM, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, CA, GN, GG, ML, MR, NE, SN, TD, TG				
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AU 2002258414	A1	20020912	AU 2002-258414	20020225
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EP 1383501	A2	20040128	EP 2002-728357	20020225
EP 1383501	B1	20070404		
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HU 200303376	A2	20040128	HU 2003-3376	20020225
JP 2004529105	T	20040924	JP 2002-567902	20020225
NZ 527364	A	20041224	NZ 2002-527364	20020225
CN 1633297	A	20050629	CN 2002-805674	20020225
BR 2002007658	A	20051025	BR 2002-7658	20020225
AT 358481	T	20070415	AT 2002-728357	20020225
US 2003225060	A1	20031204	US 2003-356879	20030203
US 6818658	B2	20041116		
ZA 2003006160	A	20040721	ZA 2003-6160	20030808
BG 108132	A	20041230	BG 2003-108132	20030825
IN 2003CN01342	A	20051125	IN 2003-CN1342	20030826
NO 2003003812	A	20031028	NO 2003-3812	20030827
US 2004266821	A1	20041230	US 2004-894719	20040720
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			US 2001-300118P	P 20010622
			WO 2002-US5724	W 20020225

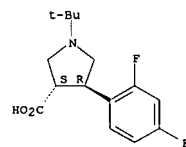
L4 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)
 US 2003-356897 A3 20030203
 OTHER SOURCE(S): MARPAT 137:216874
 GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Certain novel 4-substituted N-acylated piperidine derivs., specifically I, are agonists of the human melanocortin receptor(s) and, in particular, are selective agonists of the human melanocortin-4 receptor (MC-4R) [wherein: p = 1 or 2; q = 0, 1, or 2; n = 0, 1, or 2; R1 = H, amidino, alkyliminoyl, (un)substituted alkyl, (CH2)_n-G1 (G1 = (un)substituted cycloalkyl, Ph, naphthyl, or heteroaryl); R2 = (un)substituted Ph, naphthyl, or heteroaryl; X = alkyl, (CH2)_n-G2 (G2 = (un)substituted cycloalkyl, Ph, naphthyl, heteroaryl, heterocyclyl, cyano, CONH2, CO2H, OH, NH2, and various derivs.) where any of (CH2)_n may also be substituted; including pharmaceutically acceptable salts]. They are therefore useful for the treatment, control, or prevention of diseases and disorders responsive to the activation of MC-4R, such as obesity, diabetes, sexual dysfunction, including erectile dysfunction and female sexual dysfunction. Approx.

180 invention compds. I and approx. 25 intermediates were prepared. For instance, (2-bromo-5-chlorophenyl)acetic acid underwent a sequence of Me esterification, coupling with tert-Bu 4-[(trifluoromethyl)sulfonyloxy]-3,6-dihydropyridine-1(2H)-carboxylate via a boronate ester, removal of the BOC group, and amidation with (3S,4R)-1-(tert-butyl)-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid. The unsatd. amide-ester underwent hydrogenation, saponification of the ester, and amidation with MeNH2.HCl, to give title compound II. Representative compds. I bound to cloned human MC-4R in vitro with IC50 values generally below 2 μ M, and also acted as agonists toward cloned human MCR in a functional assay with EC50 values less than 1 μ M.
 IT 455957-75-2P
 RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
 (Intermediate; preparation of acylated piperidine derivs., particularly (pyrrolidinylcarbonyl)piperidines, (piperidinylcarbonyl)piperidines, and analogs, as melanocortin-4 receptor agonists)
 RN 455957-75-2 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, hydrochloride (1:1), (3S,4R)- (CA INDEX NAME)
 Absolute stereochemistry. Rotation (-).

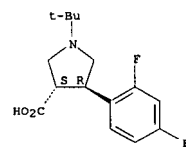
L4 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



• HCl

IT 455957-94-5, (3S,4R)-1-tert-Butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid 455957-95-6, (3R,4S)-1-tert-Butyl-4-(2,4-difluorophenyl)pyrrolidine-3-carboxylic acid
 RL: RCT (Reactant); RACT (Reactant or reagent)
 (precursor; preparation of acylated piperidine derivs., particularly (pyrrolidinylcarbonyl)piperidines, (piperidinylcarbonyl)piperidines, and analogs, as melanocortin-4 receptor agonists)
 RN 455957-94-5 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3S,4R)- (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



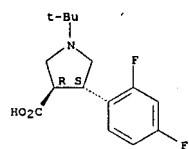
RN 455957-95-6 CAPLUS
 CN 3-Pyrrolidinecarboxylic acid, 4-(2,4-difluorophenyl)-1-(1,1-dimethylethyl)-, (3R,4S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

10/550,640

05/30/2007

L4 ANSWER 14 OF 14 CAPLUS COPYRIGHT 2007 ACS on STN (Continued)



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rxn

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LOGINID:SSPTAJMN1626

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

* * * * * Welcome to STN International * * * * *

NEWS 1 Web Page for STN Seminar Schedule - N. America
NEWS 2 JAN 08 CHEMLIST enhanced with New Zealand Inventory of Chemicals
NEWS 3 JAN 16 CA/CAPLUS Company Name Thesaurus enhanced and reloaded
NEWS 4 JAN 16 IPC version 2007.01 thesaurus available on STN
NEWS 5 JAN 16 WPIDS/WPINDEX/WPIX enhanced with IPC 8 reclassification data
NEWS 6 JAN 22 CA/CAPLUS updated with revised CAS roles
NEWS 7 JAN 22 CA/CAPLUS enhanced with patent applications from India
NEWS 8 JAN 29 PHAR reloaded with new search and display fields
NEWS 9 JAN 29 CAS Registry Number crossover limit increased to 300,000 in multiple databases
NEWS 10 FEB 15 PATDPASPC enhanced with Drug Approval numbers
NEWS 11 FEB 15 RUSSIAPAT enhanced with pre-1994 records
NEWS 12 FEB 23 KOREAPAT enhanced with IPC 8 features and functionality
NEWS 13 FEB 26 MEDLINE reloaded with enhancements
NEWS 14 FEB 26 EMBASE enhanced with Clinical Trial Number field
NEWS 15 FEB 26 TOXCENTER enhanced with reloaded MEDLINE
NEWS 16 FEB 26 IFICDB/IFIPAT/IFIUDB reloaded with enhancements
NEWS 17 FEB 26 CAS Registry Number crossover limit increased from 10,000 to 300,000 in multiple databases
NEWS 18 MAR 15 WPIDS/WPIX enhanced with new FRAGHITSTR display format
NEWS 19 MAR 16 CASREACT coverage extended
NEWS 20 MAR 20 MARPAT now updated daily
NEWS 21 MAR 22 LWPI reloaded
NEWS 22 MAR 30 RDISCLOSURE reloaded with enhancements
NEWS 23 APR 02 JICST-EPLUS removed from database clusters and STN
NEWS 24 APR 30 GENBANK reloaded and enhanced with Genome Project ID field
NEWS 25 APR 30 CHEMCATS enhanced with 1.2 million new records
NEWS 26 APR 30 CA/CAPLUS enhanced with 1870-1889 U.S. patent records
NEWS 27 APR 30 INPADOC replaced by INPADOCDB on STN
NEWS 28 MAY 01 New CAS web site launched
NEWS 29 MAY 08 CA/CAPLUS Indian patent publication number format defined
NEWS 30 MAY 14 RDISCLOSURE on STN Easy enhanced with new search and display fields
NEWS 31 MAY 21 BIOSIS reloaded and enhanced with archival data
NEWS 32 MAY 21 TOXCENTER enhanced with BIOSIS reload
NEWS 33 MAY 21 CA/CAPLUS enhanced with additional kind codes for German patents
NEWS 34 MAY 22 CA/CAPLUS enhanced with IPC reclassification in Japanese patents
NEWS EXPRESS NOVEMBER 10 CURRENT WINDOWS VERSION IS V8.01c, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 25 SEPTEMBER 2006.

NEWS HOURS STN Operating Hours Plus Help Desk Availability
NEWS LOGIN Welcome Banner and News Items
NEWS IPC8 For general information regarding STN implementation of IPC 8

Enter NEWS followed by the item number or name to see news on that specific topic.

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* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 10:47:10 ON 30 MAY 2007

=> fil casreact

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'CASREACT' ENTERED AT 10:47:23 ON 30 MAY 2007

USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT

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FILE CONTENT:1840 - 27 May 2007 VOL 146 ISS 23

New CAS Information Use Policies, enter HELP USAGETERMS for details.

* CASREACT now has more than 12 million reactions *
* *

Some CASREACT records are derived from the ZIC/VINITI database (1974-1999) provided by InfoChem, INPI data prior to 1986, and Biotransformations database compiled under the direction of Professor Dr. Klaus Kieslich.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=>

Uploading C:\Program Files\Stnexp\Queries\10550640\3.str



chain nodes :

7 8 9 10 11 12 13 14 16 17

ring nodes :

1 2 3 4 5

chain bonds :

1-8 2-7 9-10 9-12 10-11 11-16 11-17 12-13 13-14

ring bonds :

1-2 1-5 2-3 3-4 4-5

exact/norm bonds :

1-2 1-5 1-8 2-3 2-7 3-4 4-5 9-10 9-12 11-16 11-17 13-14

exact bonds :

10-11 12-13

G1:CO2H,COOH,CN

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 7:CLASS 8:CLASS 9:CLASS 10:CLASS

11:CLASS 12:CLASS 13:CLASS 14:CLASS 16:CLASS 17:CLASS

fragments assigned product role:

containing 1

fragments assigned reactant/reagent role:

containing 9

node mappings:

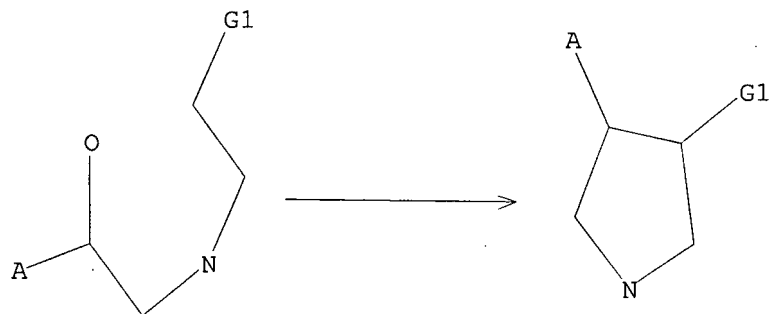
2:13 3:12 4:9 5:10 1:11

L1 STRUCTURE UPLOADED

=> d

L1 HAS NO ANSWERS

L1 STR



G1 CO2H, COOH, CN

Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 10:47:40 FILE 'CASREACT'

SCREENING COMPLETE - 6 REACTIONS TO VERIFY FROM 4 DOCUMENTS

100.0% DONE 6 VERIFIED 0 HIT RXNS 0 DOCS
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
 BATCH **COMPLETE**

PROJECTED VERIFICATIONS: 6 TO 266
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1 (0 REACTIONS)

=> s l1 full

FULL SEARCH INITIATED 10:47:47 FILE 'CASREACT'

SCREENING COMPLETE - 890 REACTIONS TO VERIFY FROM 126 DOCUMENTS

100.0% DONE 890 VERIFIED 0 HIT RXNS 0 DOCS
SEARCH TIME: 00.00.01

L3 0 SEA SSS FUL L1 (0 REACTIONS)

=>

---Logging off of STN---

=>

Executing the logoff script...